

## EAST Search History

Ref #	Hits	Search Query	DBs	Default Operator	Plurals	Time Stamp
L1	3251	"calcium citrate"	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2006/03/21 12:44
L2	13165	HDL	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2006/03/21 12:44
L3	1841	"high-density lipoprotein"	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2006/03/21 12:44
L4	13606	L2 or L3	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2006/03/21 12:45
L5	74	L1 and L4	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2006/03/21 12:49
L6	4278	postmenopausal	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2006/03/21 12:49
L7	74	L5 and L4	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2006/03/21 12:50
L8	24	L5 and L6	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2006/03/21 13:49
L9	74	L1 and L4	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2006/03/21 14:13

## EAST Search History

L10	1190	"estrogen replacement therapy"	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2006/03/21 14:14
L11	9	L7 and L10	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2006/03/21 14:18
L12	24	L6 and L9	US-PGPUB; USPAT; USOCR; EPO; JPO; DERWENT	OR	ON	2006/03/21 14:18

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID:SSSPTA1617SXX

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

\* \* \* \* \* Welcome to STN International \* \* \* \* \*

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America  
NEWS 2 "Ask CAS" for self-help around the clock  
NEWS 3 DEC 21 IPC search and display fields enhanced in CA/CAPLUS with the  
IPC reform  
NEWS 4 DEC 23 New IPC8 SEARCH, DISPLAY, and SELECT fields in USPATFULL/  
USPAT2  
NEWS 5 JAN 13 IPC 8 searching in IFIPAT, IFIUDB, and IFICDB  
NEWS 6 JAN 13 New IPC 8 SEARCH, DISPLAY, and SELECT enhancements added to  
INPADOC  
NEWS 7 JAN 17 Pre-1988 INPI data added to MARPAT  
NEWS 8 JAN 17 IPC 8 in the WPI family of databases including WPIFV  
NEWS 9 JAN 30 Saved answer limit increased  
NEWS 10 JAN 31 Monthly current-awareness alert (SDI) frequency  
added to TULSA  
NEWS 11 FEB 21 STN AnaVist, Version 1.1, lets you share your STN AnaVist  
visualization results  
NEWS 12 FEB 22 Status of current WO (PCT) information on STN  
NEWS 13 FEB 22 The IPC thesaurus added to additional patent databases on STN  
NEWS 14 FEB 22 Updates in EPFULL; IPC 8 enhancements added  
NEWS 15 FEB 27 New STN AnaVist pricing effective March 1, 2006  
NEWS 16 FEB 28 MEDLINE/LMEDLINE reload improves functionality  
NEWS 17 FEB 28 TOXCENTER reloaded with enhancements  
NEWS 18 FEB 28 REGISTRY/ZREGISTRY enhanced with more experimental spectral  
property data  
NEWS 19 MAR 01 INSPEC reloaded and enhanced  
NEWS 20 MAR 03 Updates in PATDPA; addition of IPC 8 data without attributes  
NEWS 21 MAR 08 X.25 communication option no longer available after June 2006  
  
NEWS EXPRESS FEBRUARY 15 CURRENT VERSION FOR WINDOWS IS V8.01a,  
CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),  
AND CURRENT DISCOVER FILE IS DATED 19 DECEMBER 2005.  
V8.0 AND V8.01 USERS CAN OBTAIN THE UPGRADE TO V8.01a AT  
<http://download.cas.org/express/v8.0-Discover/>  
  
NEWS HOURS STN Operating Hours Plus Help Desk Availability  
NEWS LOGIN Welcome Banner and News Items

Enter NEWS followed by the item number or name to see news on that  
specific topic.

All use of STN is subject to the provisions of the STN Customer  
agreement. Please note that this agreement limits use to scientific  
research. Use for software development or design or implementation  
of commercial gateways or other similar uses is prohibited and may  
result in loss of user privileges and other penalties.

\* \* \* \* \* STN Columbus \* \* \* \* \*

FILE 'HOME' ENTERED AT 14:32:00 ON 21 MAR 2006

=> file reg

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

0.21

0.21

FILE 'REGISTRY' ENTERED AT 14:32:42 ON 21 MAR 2006

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2006 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 20 MAR 2006 HIGHEST RN 877371-73-8

DICTIONARY FILE UPDATES: 20 MAR 2006 HIGHEST RN 877371-73-8

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH January 6, 2006

Please note that search-term pricing does apply when conducting SmartSELECT searches.

\*\*\*\*\*  
\*  
\* The CA roles and document type information have been removed from \*  
\* the IDE default display format and the ED field has been added, \*  
\* effective March 20, 2005. A new display format, IDERL, is now \*  
\* available and contains the CA role and document type information. \*  
\*  
\*\*\*\*\*

Structure search iteration limits have been increased. See HELP SLIMITS for details.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

<http://www.cas.org/ONLINE/UG/regprops.html>

=> s calcium citrate

88956 CALCIUM

5902 CITRATE

75 CITRATES

5902 CITRATE

(CITRATE OR CITRATES)

L1 6 CALCIUM CITRATE

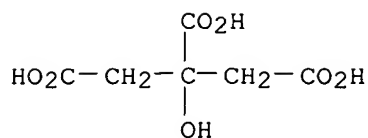
(CALCIUM(W)CITRATE)

=> s calcium citrate/cn

L2 1 CALCIUM CITRATE/CN

=> d str cn rn

L2 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2006 ACS on STN



●x Ca

\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

CN 1,2,3-Propanetricarboxylic acid, 2-hydroxy-, calcium salt (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN **Calcium citrate (6CI)**

CN Citric acid, calcium salt (8CI)

OTHER NAMES:

CN Citramar

CN E 333

RN 7693-13-2 REGISTRY

=> file caplus medline biosis embase

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

17.06

17.27

FILE 'CAPLUS' ENTERED AT 14:33:46 ON 21 MAR 2006

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'MEDLINE' ENTERED AT 14:33:46 ON 21 MAR 2006

FILE 'BIOSIS' ENTERED AT 14:33:46 ON 21 MAR 2006

Copyright (c) 2006 The Thomson Corporation

FILE 'EMBASE' ENTERED AT 14:33:46 ON 21 MAR 2006

Copyright (c) 2006 Elsevier B.V. All rights reserved.

=> s 7693-13-2/rn

'RN' IS NOT A VALID FIELD CODE

'RN' IS NOT A VALID FIELD CODE

'RN' IS NOT A VALID FIELD CODE

L3 1162 7693-13-2/RN

=> s calcium citrate

L4 2063 CALCIUM CITRATE

=> s calcium and citrate

L5 17375 CALCIUM AND CITRATE

=> s L4 or L5

L6 17375 L4 OR L5

=> s L3 or L6

L7 17512 L3 OR L6

=> s HDL or high density lipoprotein

L8 112539 HDL OR HIGH DENSITY LIPOPROTEIN

=> s HDL or high density lipoprotein  
L9 150985 HDL OR HIGH DENSITY LIPOPROTEIN

=> s high density lipoprotein  
L10 102670 HIGH DENSITY LIPOPROTEIN

=> s L7 and L8  
L11 17 L7 AND L8

=> dup rem L11  
PROCESSING COMPLETED FOR L11  
L12 11 DUP REM L11 (6 DUPLICATES REMOVED)

=> s postmenopausal  
L13 82255 POSTMENOPAUSAL

=> s L12 and L13  
L14 3 L12 AND L13

=> d 1-3 ibib abs

L14 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER: 2002:696671 CAPLUS  
DOCUMENT NUMBER: 137:216323  
TITLE: Method of administering **calcium citrate**  
INVENTOR(S): Reid, Ian R.  
PATENT ASSIGNEE(S): Uniservices Ltd., N. Z.  
SOURCE: U.S. Pat. Appl. Publ., 13 pp.  
CODEN: USXXCO  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002128320	A1	20020912	US 2001-16371	20011210
WO 2003049668	A2	20030619	WO 2002-IB5759	20021210
WO 2003049668	A3	20040617		

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW  
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.: US 2001-16371 A 20011210

AB A method of increasing a high-d. lipoprotein level in plasma of a **postmenopausal** woman by administering a pharmaceutical formulation containing **calcium citrate** is described. The therapeutically ED of **calcium citrate** is equivalent to at least about 1 g elemental **calcium**. An oral pharmaceutical composition and a dietary supplement comprises **calcium citrate** in an amount sufficient to provide about 10 mg to about 1 g elemental **calcium** to a diet of a **postmenopausal** woman.

L14 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER: 2002:211581 CAPLUS  
DOCUMENT NUMBER: 136:385392  
TITLE: Effects of **calcium** supplementation on serum

lipid concentrations in normal older women: A  
randomized controlled trial  
AUTHOR(S): Reid, Ian R.; Mason, Barbara; Horne, Anne; Ames, Ruth;  
Clearwater, Judith; Bava, Usha; Orr-Walker, Brandon;  
Wu, Fiona; Evans, Margaret C.; Gamble, Gregory D.  
CORPORATE SOURCE: Department of Medicine, University of Auckland,  
Auckland, N. Z.  
SOURCE: American Journal of Medicine (2002), 112(5), 343-347  
CODEN: AJMEAZ; ISSN: 0002-9343  
PUBLISHER: Excerpta Medica, Inc.  
DOCUMENT TYPE: Journal  
LANGUAGE: English

AB To determine the effect of supplementation with Ca **citrate** on  
circulating lipid concns. in normal older women. As part of a study of  
the effects of Ca supplementation on fractures, the authors randomly  
assigned 223 **postmenopausal** women (mean [ $\pm$  SD] age, 72  $\pm$  4  
yr), who were not receiving therapy for hyperlipidemia or osteoporosis, to  
receive Ca (1 g/d, n = 111) or placebo (n = 112) for 1 yr. Fasting serum  
lipid concns., including high-d. lipoprotein (HDL) cholesterol  
and low-d. lipoprotein (LDL) cholesterol, were obtained at baseline, and  
at 2, 6, and 12 mo. After 12 mo, HDL cholesterol levels and the  
HDL cholesterol to LDL cholesterol ratio had increased more in the  
Ca group than in the placebo group (mean between-group differences in  
change from baseline: for HDL cholesterol, 0.09 mmol/L (95%  
confidence interval [CI]: 0.02 to 0.17; P = 0.01); for HDL/LDL  
cholesterol ratio, 0.05 (95% CI: 0.02 to 0.08; P = 0.001)). This was  
largely due to a 7% increase in HDL cholesterol levels in the Ca  
group, with a nonsignificant 6% decline in LDL cholesterol levels. There  
was no significant treatment effect on triglyceride level (P = 0.48).  
**Calcium citrate** supplementation causes beneficial  
changes in circulating lipids in **postmenopausal** women. This  
suggests that a reappraisal of the indications for Ca supplementation is  
necessary, and that its cost effectiveness may were underestimated.

REFERENCE COUNT: 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 3 OF 3 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights  
reserved on STN

ACCESSION NUMBER: 2004070368 EMBASE  
TITLE: Effects of **Calcium** Supplementation on Circulating  
Lipids: Potential Pharmacoeconomic Implications.  
AUTHOR: Reid I.R.  
CORPORATE SOURCE: Prof. I.R. Reid, Department of Medicine, University of  
Auckland, Private Bag 92019, Auckland, New Zealand.  
i.reid@auckland.ac.nz  
SOURCE: Drugs and Aging, (2004) Vol. 21, No. 1, pp. 7-17. .  
Refs: 34  
ISSN: 1170-229X CODEN: DRAGE6  
COUNTRY: New Zealand  
DOCUMENT TYPE: Journal; General Review  
FILE SEGMENT: 018 Cardiovascular Diseases and Cardiovascular Surgery  
036 Health Policy, Economics and Management  
037 Drug Literature Index  
038 Adverse Reactions Titles  
048 Gastroenterology  
LANGUAGE: English  
SUMMARY LANGUAGE: English  
ENTRY DATE: Entered STN: 20040304  
Last Updated on STN: 20040304

AB For about a century there has been recognition that **calcium** and  
lipids bind to one another in the gut, each interfering with the other's  
absorption. **Calcium** also causes malabsorption of bile acids,  
which is likely to contribute further to malabsorption of fat. High  
dietary **calcium** intakes may also have stimulatory effects on

lipolysis. These mechanisms provide a basis for hypothesising that **calcium** supplementation may impact on circulating lipid concentrations, and there is now a significant amount of observational and trial data indicating that this is the case. The largest randomised controlled trial of **calcium** effects on lipids was carried out in 223 healthy **postmenopausal** women, and found that low density lipoprotein-cholesterol (LDL-C) decreased 6.3% and high density lipoprotein-cholesterol (HDL-C) increased by 7.3% at 1-year. The resultant 16.4% increase in HDL-C/LDL-C ratio would be predicted to reduce cardiovascular event rates by 20-30%, which is consistent with the available observational data. There are no trial data addressing this question and it is possible that other lipid-lowering agents, such as hydroxymethylglutaryl coenzyme A reductase inhibitors, might impact on cardiac event rates by mechanisms other than by lowering cholesterol levels. Therefore, caution is appropriate in incorporating these findings into clinical practice, but the balance of evidence suggests that **calcium** is a cost-effective adjunct to the dietary management of hyperlipidaemia.

=> d 1-11 L12 ibib abs

L12 ANSWER 1 OF 11 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2005:411059 CAPLUS  
 DOCUMENT NUMBER: 142:469260  
 TITLE: HDL-boosting combination therapy complexes  
 INVENTOR(S): Tunac, Josefino B.  
 PATENT ASSIGNEE(S): USA  
 SOURCE: U.S. Pat. Appl. Publ., 15 pp.  
 CODEN: USXXCO  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2005101561	A1	20050512	US 2004-983836	20041108
WO 2005046662	A2	20050526	WO 2004-US37324	20041108
WO 2005046662	A3	20050623		

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.: US 2003-518091P P 20031107

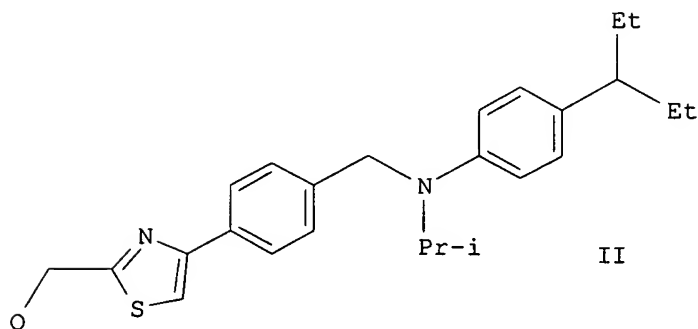
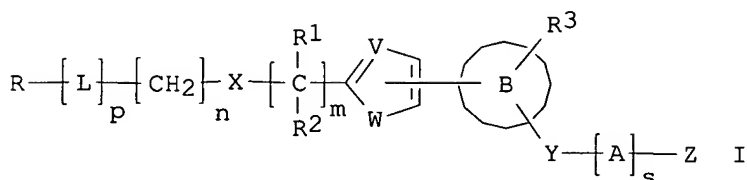
AB A pharmaceutical composition including therapeutically effective amts. of at least one HMG-CoA reductase inhibitor present as a dyhydroxyacid salt and at least one addnl. therapeutic agent is claimed. Dehydroxy acid salt of sodium lovastatin (I) was prepared and its antilipidemic activity was studied in hamster. A repeat side-by-side comparison between I and Lipitor at 5-20 mg dose range confirmed the effectiveness of I in decreasing LDL, moreover, I was effective at a dose as low as 5 mg.

L12 ANSWER 2 OF 11 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2004:878382 CAPLUS  
 DOCUMENT NUMBER: 141:350161  
 TITLE: Preparation of azole compounds as PTP1B inhibitors



INVENTOR(S): Ikemoto, Tomoyuki; Tanaka, Masahiro; Yuno, Takeo;  
Sakamoto, Johei; Nakanishi, Hiroyuki; Nakagawa,  
Yuichi; Ohta, Takeshi; Sakata, Shohei; Morinaga,  
Hisayo  
PATENT ASSIGNEE(S): Japan Tobacco Inc., Japan  
SOURCE: PCT Int. Appl., 542 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: Japanese  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004089918	A1	20041021	WO 2004-JP5119	20040409
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CA 2521830	AA	20041021	CA 2004-2521830	20040409
EP 1553091	A1	20050713	EP 2004-726765	20040409
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR				
JP 2005272476	A2	20051006	JP 2005-133755	20050428
PRIORITY APPLN. INFO.:			JP 2003-105267	A 20030409
			JP 2003-157590	A 20030603
			JP 2005-505323	A3 20040409
			WO 2004-JP5119	W 20040409
OTHER SOURCE(S):	MARPAT 141:350161			
GI				



AB Title compds. I [V = N, CH; W = S, O; m = 0-2; R1, R2 = H, alkyl; X = NR4, etc.; R4 = H, alkyl; n = 0-4; p = 0, 1; L = CR2OR21, etc.; R20 = H, alkyl, etc.; R21 = H, alkyl, etc.; R = CO2R19, etc.; R19 = H, alkyl; B = aryl, heteroaryl; R3 = H, halo, etc.; Y = O, etc.; s = 0, 1; A = (un)substituted alkylene with cycloalkyl; Z = cycloalkyl, etc.] were prepared For example, O-alkylation of 5-hydroxynicotinic acid Me ester with compound II [Q = Cl], e.g., prepared from 4-bromoacetylbenzoic acid in 5 steps, followed by

saponification

afforded compound II [3-carboxypyridin-5-yloxy] in 44.1% overall yield. In PTP1B (protein tyrosine phosphatase 1B) inhibition assays, the IC50 value of compound II [Q = 3-carboxypyridin-5-yloxy] was 0.28  $\mu$ M. Compds. I are claimed useful for the treatment of obesity, diabetes, etc. Formulations are given.

REFERENCE COUNT: 16 THERE ARE 16 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 3 OF 11 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights reserved on STN

ACCESSION NUMBER: 2004070368 EMBASE

TITLE: Effects of **Calcium** Supplementation on Circulating Lipids: Potential Pharmacoeconomic Implications.

AUTHOR: Reid I.R.

CORPORATE SOURCE: Prof. I.R. Reid, Department of Medicine, University of Auckland, Private Bag 92019, Auckland, New Zealand. i.reid@auckland.ac.nz

SOURCE: Drugs and Aging, (2004) Vol. 21, No. 1, pp. 7-17. . Refs: 34

ISSN: 1170-229X CODEN: DRAGE6

COUNTRY: New Zealand

DOCUMENT TYPE: Journal; General Review

FILE SEGMENT: 018 Cardiovascular Diseases and Cardiovascular Surgery  
036 Health Policy, Economics and Management  
037 Drug Literature Index  
038 Adverse Reactions Titles  
048 Gastroenterology

LANGUAGE: English

SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 20040304

Last Updated on STN: 20040304

AB For about a century there has been recognition that **calcium** and lipids bind to one another in the gut, each interfering with the other's absorption. **Calcium** also causes malabsorption of bile acids, which is likely to contribute further to malabsorption of fat. High dietary **calcium** intakes may also have stimulatory effects on lipolysis. These mechanisms provide a basis for hypothesising that **calcium** supplementation may impact on circulating lipid concentrations, and there is now a significant amount of observational and trial data indicating that this is the case. The largest randomised controlled trial of **calcium** effects on lipids was carried out in 223 healthy postmenopausal women, and found that low density lipoprotein-cholesterol (LDL-C) decreased 6.3% and high density lipoprotein-cholesterol (HDL-C) increased by 7.3% at 1-year. The resultant 16.4% increase in HDL-C/LDL-C ratio would be predicted to reduce cardiovascular event rates by 20-30%, which is consistent with the available observational data. There are no trial data addressing this question and it is possible that other lipid-lowering agents, such as hydroxymethylglutaryl coenzyme A reductase inhibitors, might impact on cardiac event rates by mechanisms other than by lowering cholesterol levels. Therefore, caution is appropriate in incorporating these findings into clinical practice, but the balance of evidence suggests that **calcium** is a cost-effective adjunct to the dietary management of hyperlipidaemia.

L12 ANSWER 4 OF 11 MEDLINE on STN  
 ACCESSION NUMBER: 2002342613 MEDLINE  
 DOCUMENT NUMBER: PubMed ID: 12085783  
 TITLE: **Calcium's** healthy cholesterol consequences.  
 AUTHOR: Anonymous  
 SOURCE: Health news (Waltham, Mass.), (2002 Jun) Vol. 8, No. 6, pp. 5.  
 Journal code: 9800495. ISSN: 1081-5880.  
 PUB. COUNTRY: United States  
 DOCUMENT TYPE: News Announcement  
 LANGUAGE: English  
 FILE SEGMENT: Consumer Health  
 ENTRY MONTH: 200206  
 ENTRY DATE: Entered STN: 20020628  
 Last Updated on STN: 20020628  
 Entered Medline: 20020627

L12 ANSWER 5 OF 11 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2002:696671 CAPLUS  
 DOCUMENT NUMBER: 137:216323  
 TITLE: Method of administering **calcium citrate**  
 INVENTOR(S): Reid, Ian R.  
 PATENT ASSIGNEE(S): Uniservices Ltd., N. Z.  
 SOURCE: U.S. Pat. Appl. Publ., 13 pp.  
 CODEN: USXXCO  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002128320	A1	20020912	US 2001-16371	20011210
WO 2003049668	A2	20030619	WO 2002-IB5759	20021210
WO 2003049668	A3	20040617		

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW  
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.: US 2001-16371 A 20011210  
 AB A method of increasing a high-d. lipoprotein level in plasma of a postmenopausal woman by administering a pharmaceutical formulation containing **calcium citrate** is described. The therapeutically ED of **calcium citrate** is equivalent to at least about 1 g elemental **calcium**. An oral pharmaceutical composition and a dietary supplement comprises **calcium citrate** in an amount sufficient to provide about 10 mg to about 1 g elemental **calcium** to a diet of a postmenopausal woman.

L12 ANSWER 6 OF 11 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2002:392237 CAPLUS  
 DOCUMENT NUMBER: 136:401651  
 TITLE: Preparation of fused pyridine derivatives as HMG-CoA reductase inhibitors  
 INVENTOR(S): Robl, Jeffrey A.; Chen, Bang-Chi; Sun, Chong-Qing  
 PATENT ASSIGNEE(S): USA  
 SOURCE: U.S. Pat. Appl. Publ., 46 pp., Cont.-in-part of U.S.

Ser. No. 875,218.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

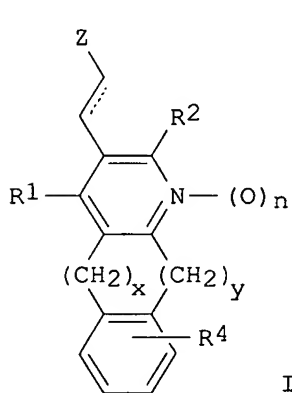
FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

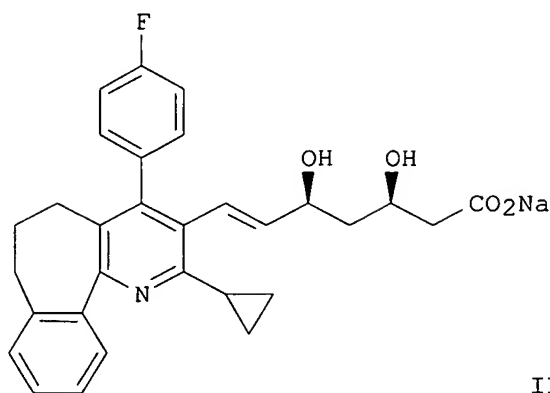
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002061901	A1	20020523	US 2001-8154	20011204
US 6620821	B2	20030916		
US 2002028826	A1	20020307	US 2001-875218	20010606
US 2004024216	A1	20040205	US 2003-602753	20030624
PRIORITY APPLN. INFO.:			US 2000-211594P	P 20000615
			US 2001-875218	A2 20010606
			US 2001-8154	A3 20011204

OTHER SOURCE(S): MARPAT 136:401651

GI



I



II

AB The title compds. I and their pharmaceutically acceptable salts, esters, prodrug esters, and stereoisomers are claimed [wherein: Z = CH(OH)CH<sub>2</sub>CR<sub>7</sub>(OH)CH<sub>2</sub>CO<sub>2</sub>R<sub>3</sub> or corresponding pyranone lactone derivs.; n = 0, 1; x = 0, 1, 2, 3, or 4; y = 0, 1, 2, 3 or 4, provided that at least one of x and y is other than 0; and optionally one or more carbons of (CH<sub>2</sub>)<sub>x</sub> and/or (CH<sub>2</sub>)<sub>y</sub> together with addnl. carbons form a 3 to 7 membered spirocyclic ring; R<sub>1</sub>, R<sub>2</sub> = alkyl, arylalkyl, cycloalkyl, alkenyl, cycloalkenyl, aryl, heteroaryl, cycloheteroalkyl; R<sub>3</sub> = H or lower alkyl; R<sub>4</sub> = H, halo, CF<sub>3</sub>, OH, alkyl, alkoxy, CO<sub>2</sub>H, (un)substituted NH<sub>2</sub>, cyano, (un)substituted CONH<sub>2</sub>, etc.; R<sub>7</sub> = H, alkyl]. The compds. are HMG-CoA reductase inhibitors, and are active in inhibiting cholesterol biosynthesis and modulating blood serum lipids, for example, lowering LDL cholesterol and/or increasing HDL cholesterol (no data). I are thus useful in treating hyperlipidemia and dyslipidemia, in hormone replacement therapy, and in treating hypercholesterolemia, hypertriglyceridemia and atherosclerosis, as well as Alzheimer's disease and osteoporosis. Prepns. of several compds. are described. For instance, a multistep synthesis of fused pyridine derivative II is reported. Compds. I may be used in a manner similar to atorvastatin, pravastatin, simvastatin, etc. Combinations of compds. I with various other drugs are claimed, the latter being specified as certain pharmacol. classes, as inhibitors of specific enzymes, as (ant)agonists of specific receptors, and as numerous named drugs.

L12 ANSWER 7 OF 11 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 1

ACCESSION NUMBER: 2002:211581 CAPLUS

DOCUMENT NUMBER: 136:385392

TITLE: Effects of **calcium** supplementation on serum lipid concentrations in normal older women: A randomized controlled trial

AUTHOR(S): Reid, Ian R.; Mason, Barbara; Horne, Anne; Ames, Ruth; Clearwater, Judith; Bava, Usha; Orr-Walker, Brandon; Wu, Fiona; Evans, Margaret C.; Gamble, Gregory D.

CORPORATE SOURCE: Department of Medicine, University of Auckland, Auckland, N. Z.

SOURCE: American Journal of Medicine (2002), 112(5), 343-347  
CODEN: AJMEAZ; ISSN: 0002-9343

PUBLISHER: Excerpta Medica, Inc.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB To determine the effect of supplementation with Ca **citrate** on circulating lipid concns. in normal older women. As part of a study of the effects of Ca supplementation on fractures, the authors randomly assigned 223 postmenopausal women (mean [ $\pm$  SD] age, 72  $\pm$  4 yr), who were not receiving therapy for hyperlipidemia or osteoporosis, to receive Ca (1 g/d, n = 111) or placebo (n = 112) for 1 yr. Fasting serum lipid concns., including high-d. lipoprotein (**HDL**) cholesterol and low-d. lipoprotein (LDL) cholesterol, were obtained at baseline, and at 2, 6, and 12 mo. After 12 mo, **HDL** cholesterol levels and the **HDL** cholesterol to LDL cholesterol ratio had increased more in the Ca group than in the placebo group (mean between-group differences in change from baseline: for **HDL** cholesterol, 0.09 mmol/L (95% confidence interval [CI]: 0.02 to 0.17; P = 0.01); for **HDL**/LDL cholesterol ratio, 0.05 (95% CI: 0.02 to 0.08; P = 0.001)). This was largely due to a 7% increase in **HDL** cholesterol levels in the Ca group, with a nonsignificant 6% decline in LDL cholesterol levels. There was no significant treatment effect on triglyceride level (P = 0.48). **Calcium citrate** supplementation causes beneficial changes in circulating lipids in postmenopausal women. This suggests that a reappraisal of the indications for Ca supplementation is necessary, and that its cost effectiveness may have been underestimated.

REFERENCE COUNT: 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 8 OF 11 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2001:747110 CAPLUS

DOCUMENT NUMBER: 135:256481

TITLE: Manufacture of a cultured dairy product containing exogenously added protein

INVENTOR(S): Nadland, Karl Johan

PATENT ASSIGNEE(S): Nutri Pharma Asa, Norway

SOURCE: Eur. Pat. Appl., 22 pp.  
CODEN: EPXXDW

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
-----	----	-----	-----	-----
EP 1142482	A1	20011010	EP 2000-610033	20000404
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
WO 2001074171	A1	20011011	WO 2001-IB553	20010403
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,  
DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,  
BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

AU 2001046748 A5 20011015 AU 2001-46748 20010403

US 2002012719 A1 20020131 US 2001-828486 20010409

PRIORITY APPLN. INFO.:

EP 2000-610033 A 20000404

US 2000-195988P P 20000407

WO 2001-IB553 W 20010403

AB Cultured dairy products are formulated to contain exogenously added protein and optionally exogenously added dietary fiber. The methods comprise the steps of (i) hydrating a protein source by subjecting it to shear forces and if necessary to heat in the presence of excess of water; (ii) adding the hydrated protein source from step (i) to a milk composition; (iii) adding a fermentation culture to the mixture from step (ii); and (iv) fermenting to obtain a cultured dairy product. The shear forces are preferably applied by use of a homogenizer at a temperature of 80°. The exogenously added proteins are preferably soy proteins and the exogenously added dietary fibers are preferably soybean fiber, especially soybean cotyledon fibers. The cultured dairy products preferably contain exogenously added protein in an amount of  $\geq 5\%$  by weight. Thus, a suitable protein source may include soy protein 10.66, soybean cotyledon fiber 2.67, and soy lecithin 0.76% (weight percent of cultured dairy product).

REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 9 OF 11 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights reserved on STN

ACCESSION NUMBER: 2000398596 EMBASE

TITLE: Heparin-free DALI LDL-apheresis in hyperlipidemic patients: Efficacy, safety and biocompatibility.

AUTHOR: Wendler T.; Lennertz A.; Heinemann O.; Duhr C.; Samtleben W.; Bosch T.

CORPORATE SOURCE: Dr. T. Bosch, Schwerpunkt Nephrologie, Medizinische Klinik I, Klin. Grosshadern der Univ. Munchen, D-81366 Munchen, Germany. bosch@med1.uni-muenchen.de

SOURCE: International Journal of Artificial Organs, (2000) Vol. 23, No. 10, pp. 710-717. .  
Refs: 19

ISSN: 0391-3988 CODEN: IJAODS

COUNTRY: Italy

DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 018 Cardiovascular Diseases and Cardiovascular Surgery  
006 Internal Medicine  
030 Pharmacology  
025 Hematology  
029 Clinical Biochemistry  
038 Adverse Reactions Titles  
037 Drug Literature Index  
003 Endocrinology

LANGUAGE: English

SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 20001213

Last Updated on STN: 20001213

AB Background and aim of the study. In routine DALI apheresis - the first technique for direct adsorption of lipoproteins from whole blood - heparin plus **citrate** (ACD-A) is used as anticoagulation regimen. However, recently several publications have warned of heparin-induced thrombocytopenia as a rare but potentially life-threatening complication of heparin administration (HIT type 2). The aim of the present study was therefore to test the efficacy and biocompatibility of DALI using a heparin-free anticoagulation regimen consisting exclusively of **citrate**. Methods. Four symptomatic hypercholesterolemic patients on regular DALI apheresis were switched to the heparin-free protocol for two sessions each. Two of the patients were on oral anticoagulation using

phenprocoumon. In the weekly sessions, 1.3 patient blood volumes were processed at a blood flow rate of 60 ml/min using ACD-A at a ratio of 1:20 (v/v) during adsorber priming and the session. Results. Clinically, all sessions were essentially uneventful. Uncorrected lipoprotein reductions amounted to 65% for LDL-C, 62% for Lp(a), 53% for VLDL-C, 24% for HDL-C, 17% for triglycerides and 19% for fibrinogen. Cell counts remained virtually constant. No signs of hemolysis or clotting could be detected. Thromboplastin time (Quick) was slightly prolonged and partial thromboplastin time (PTT) moderately elevated in all patients. In contrast, whole blood coagulation time acc. to Lee-White and activated clotting times were increased only in orally anticoagulated patients. Biocompatibility in terms of complement, leukocyte and thrombocyte activation was excellent. Bradykinin activation was moderate peaking at 3038 pg/ml in the efferent line. Systemic thrombin-antithrombin complex (TAT) reflected perfect anticoagulation in orally anticoagulated patients and adequate anticoagulation in the patients without phenprocoumon. Conclusion. In this pilot study heparin-free DALI apheresis was safe and effective and may thus be performed in LDL-apheresis dependent patients who suffer from heparin intolerance.

L12 ANSWER 10 OF 11 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 2

ACCESSION NUMBER: 1993:471434 CAPLUS

DOCUMENT NUMBER: 119:71434

TITLE: Short-term dietary **calcium** fortification increases fecal saturated fat content and reduces serum lipids in men

AUTHOR(S): Denke, Margo A.; Fox, Mary M.; Schulte, Marcia C.

CORPORATE SOURCE: Southwest. Med. Cent., Univ. Texas, Dallas, TX, 75235-9052, USA

SOURCE: Journal of Nutrition (1993), 123(6), 1047-53

CODEN: JONUAI; ISSN: 0022-3166

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The effect of dietary Ca on fecal fatty acid excretion and serum lipids was tested in a randomized, single-blind metabolic study in 13 healthy men with moderate hypercholesterolemia. A low-Ca base diet containing 34% of energy from fat, 13% from saturated fatty acids, 240 mg cholesterol/day, and 410 mg Ca/day was compared with a fortified version in which Ca **citrate** malate was added to orange juice (550 mg), muffins (750 mg), and 2 tablets (500 mg) for a total Ca intake of 2200 mg/day. Fecal collections (72 h, days 8, 9, 10) and blood from fasting subjects for lipids and lipoproteins (days 9, 10, 11) were obtained. The percentage of dietary saturated fat excreted per day increased from 6 to 13% with Ca fortification. There was no change in fecal bile acid excretion. The high-Ca diet significantly reduced total cholesterol by 6% (5.99 to 5.66 mmol/L), LDL cholesterol by 11% (4.13 to 3.67 mmol/L), and apolipoprotein B concns. by 7% when compared with the low-Ca diet. There was no change in HDL cholesterol or apolipoprotein A1 concns. Urinary Ca excretion increased from 146 to 230 mg/day when the high-Ca diet was consumed. Ca fortification was effective in lowering total and LDL cholesterol concns. and may be an effective adjunct to cholesterol-lowering diet therapy.

L12 ANSWER 11 OF 11 MEDLINE on STN

ACCESSION NUMBER: 87049045 MEDLINE

DOCUMENT NUMBER: PubMed ID: 3778574

TITLE: Atherogenesis. Mitigation of monocyte adhesion to arterial endothelium in hyperlipidemic rats by phosphocitrate, a phosphorylated polycarboxylic acid.

AUTHOR: Shankar R; Tuyethong N; Sallis J D

SOURCE: Atherosclerosis, (1986 Oct) Vol. 62, No. 1, pp. 47-54.

Journal code: 0242543. ISSN: 0021-9150.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 198612  
ENTRY DATE: Entered STN: 19900302  
Last Updated on STN: 19900302  
Entered Medline: 19861215

AB Phosphocitrate, a phosphorylated polycarboxylic acid ameliorates two early events in atherogenesis. When administered to rats on an atherosclerotic diet (112 mumol/kg body weight/day), it reduced monocyte adhesion to aortic endothelium from 34 +/- 7 cells/HPF for untreated rats to 1 +/- 1 cell/HPF, a value seen in normal, non-atherosclerotic rats. Transmission electron microscopy of aortic sections showed no evidence of subendothelial lipid accumulation in phosphocitrate-treated rats despite the high circulating plasma lipid levels. The mechanisms of action of phosphocitrate are unknown but the indications are that its influence may be mediated through its polyanionic chemical nature and/or its ability to modulate cellular **calcium** accumulation. In addition to its possible therapeutic value as an anti-calcifying and anti-atherogenic compound, phosphocitrate may prove useful as an experimental probe for studying the cellular basis of atherogenesis.

=> FIL STNGUIDE  
COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
84.79	102.06

FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE	TOTAL
ENTRY	SESSION
-6.75	-6.75

CA SUBSCRIBER PRICE

FILE 'STNGUIDE' ENTERED AT 14:46:11 ON 21 MAR 2006  
USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT  
COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY, JAPAN SCIENCE  
AND TECHNOLOGY CORPORATION, AND FACHINFORMATIONSZENTRUM KARLSRUHE

FILE CONTAINS CURRENT INFORMATION.  
LAST RELOADED: Mar 17, 2006 (20060317/UP).